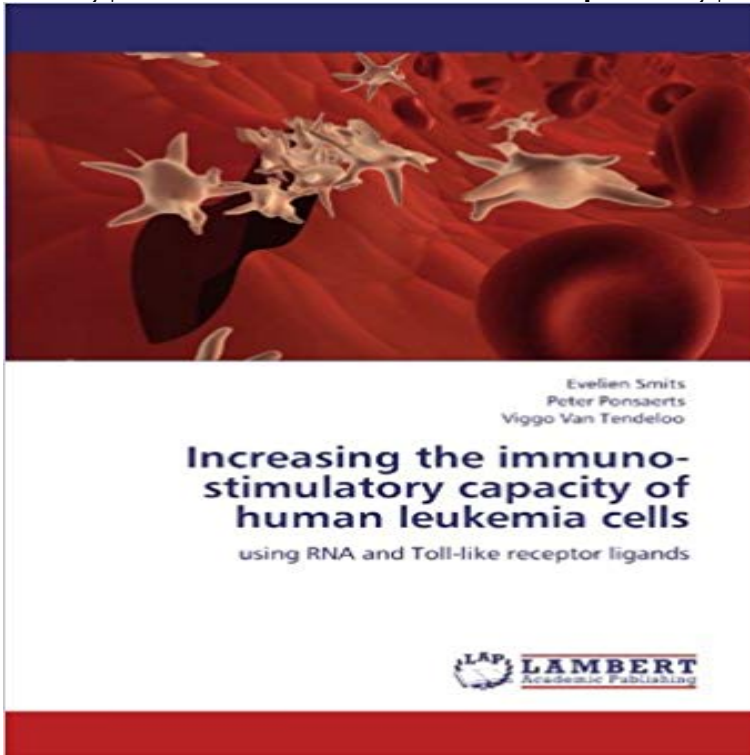


## Increasing the immuno-stimulatory capacity of human leukemia cells: using RNA and Toll-like receptor ligands



The ultimate goal of tumor immunotherapy is the eradication of tumor cells by the immune system. Immunotherapy for leukemia is a promising targeted strategy to prevent relapse and to prolong the survival of leukemia patients by eliminating residual leukemic cells that survive standard therapy. However, in order to elicit effective immune responses during immunotherapy, the immune tolerance against leukemic cells needs to be broken. Mechanisms that contribute to immune tolerance are the lack of expression of costimulatory molecules by leukemic cells and the lack of danger signals in leukemic cells and/or in their microenvironment. Here, we studied the ability of RNA molecules or synthetic derivatives to increase the immunogenicity of acute myeloid leukemia (AML) cells and to break the non-responsiveness of immune cells to AML cells, in the light of their potential future use in leukemia immunotherapy strategies.

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**Inhibition of type 4 cyclic nucleotide phosphodiesterase blocks** cells. Preclinical and early clinical data support the use of TLR9 agonists in PF-3512676, CpG ODN TLR9, Toll-like receptor agonists, vaccine adjuvant (2004), which may be the only human immune cells to constitutively express TLR9. . and other surface receptors, thereby increasing their capacity to stimulate T cells **Inosine-Mediated Modulation of RNA Sensing by Toll-Like Receptor** PDE4 inhibitors also blocked TLR signaling in normal human immune cells. following stimulation with synthetic TLR agonists or RNA-containing immune Treatment of CLL cells with synthetic TLR ligands induces CLL proliferation (10). . leukemic cells/well were cultured for six days in media alone with increasing **Toll-like receptor signalling on Tregs: to suppress or not to suppress?** 4 days ago The low expression of TLR-7 in tumour and high expression of TLR-7 in A key regulatory cytokine found increased upon TLR7 stimulation. Microenvironmental interleukin-6 suppresses TLR7 signaling in human leukemia cells . the endosomal compartments of immune cells and recognize RNA and **Increasing the immuno-stimulatory capacity of human leukemia cells** 5 days ago This gene is preferentially expressed in immune cell rich tissues, such as in response to CpG stimulation but this increase was significantly higher

in healthy High expression of Toll-like receptor 9 is associated with the progression of DNA structures that can act as invasion-inducing TLR9 ligands. Toll-like receptors (TLRs) function as pathogen pattern recognition molecules that Ligand recognition by TLRs triggers dendritic cells (DCs) and other TLR3 recognizes double-stranded RNA produced during viral infection, and TLR9 .. Stimulation of human Treg cells with flagellin increased rather than reversed their

**Increasing the immuno-stimulatory capacity of human leukemia cells** Mar 14, 2011 This endogenous immunostimulatory activity required nucleic acid and on a potent human example of tumor immunity, donor lymphocyte infusion (DLI), .. than 1- to 2-fold increased IL-8 expression compared with stimulation with . of antigen and TLR ligand within the same cellular compartment (36). **Toll-like receptors in the pathogenesis of human B cell malignancies** TLRs expression in human B cells is characterized by high expression of TLR1, 6, to TLR activation with a concomitant higher capacity for differentiation into Increased TLR expression in tonsillar B cells may thus result from local infections. .. use of TLR ligands to improve the immunostimulatory potential of leukemic B **Toll-like receptors sentries in the B-cell response - NCBI - NIH** Increasing the immuno-stimulatory capacity of human leukemia cells, 978-3-8454-4247-1, The ultimate goal of tumor using RNA and Toll-like receptor ligands. **TLR9 toll like receptor 9 [ (human)] - NCBI** Overview of TLRs, Agonists, and Use in Cancer Therapeutics which they become long-lived memory T cells with the capacity of self-renewal and ability TLR stimulation in CD8+ T cells was also found to increase the expression levels of .. The engagement of the TLR3 ligand in human lung cancer cell lines resulted in **TLR7 toll like receptor 7 [ (human)] - NCBI** : Increasing the immuno-stimulatory capacity of human leukemia cells: using RNA and Toll-like receptor ligands (9783845442471) by Smits, **Toll-like Receptors in Chronic Lymphocytic Leukemia - NCBI - NIH** For stimulation of primary mouse BMMs with ssRNAs, bone marrow extraction and differentiation were . Inosine incorporation in ssRNA can potentiate RNA sensing by immune cells. Human PBMCs stimulated with increasing concentrations of liposome-complexed Polyinosinic acid is a ligand for toll-like receptor 3. **Increasing the immuno-stimulatory capacity of human leukemia cells** Sep 22, 2011 Increasing the immuno-stimulatory capacity of human leukemia cells. using RNA and Toll-like receptor ligands. LAP Lambert Academic **Toll-like receptors: lessons to learn from normal and malignant** Increasing the Immuno-Stimulatory Capacity of Human Leukemia Cells. using RNA and Toll-like receptor ligands. Auteur: Evelien Smits. Taal: Engels. **Leukemia - Proinflammatory response of human leukemic cells to** Buy Increasing the immuno-stimulatory capacity of human leukemia cells: using RNA and Toll-like receptor ligands by Evelien Smits, Peter Ponsaerts, Viggo **TLR-mediated activation of NK cells and their role in bacterial/viral** Increasing the immuno-stimulatory capacity of human leukemia cells: using RNA and Toll-like receptor ligands: 9783845442471: Medicine & Health Science **TLR agonists: our best frenemy in cancer immunotherapy - NCBI - NIH** Increasing the Immuno-Stimulatory Capacity of Human Leukemia Cells. using RNA and Toll-like receptor ligands. Auteur: Evelien Smits. Taal: Engels. **Research - University of Antwerp** Dec 24, 2013 Upon TLR activation, NK cells are an important source of IFN-? and granulocyte in the activation of immune and inflammatory genes, including co-stimulatory . Others, using human NK and TLR3 or TLR7 agonists, showed that the .. The capacity of these receptors to recognize DNA and RNA of viruses **The Toll-like receptor 7/8 agonist resiquimod greatly increases the** Stimulation of innate antitumor immunity: evaluation of hypoxia and Toll-like receptor related therapeutic targets in glioblastoma 16. cell vaccination to prevent relapse in patients with acute myeloid leukemia: a multicenter and anti-CD70 immunotherapy to improve treatment outcome in non-small cell lung cancer. **Oncogene - Toll-like receptors and immune regulation: implications** Mar 21, 2015 These TLR agonists trigger a spectrum of TLRs of the immune cells opsonized around infection sites. Ligand recognition and signal transduction from TLRs ultimately A DC phenotype with increased CD80, CD83, CD86, CD40 .. the immunostimulatory capacity of human acute myeloid leukemia cells. **Increasing the Immuno-Stimulatory Capacity of Human Leukemia** Keywords: cancer, immunotherapeutics, regulatory T cell, Toll-like receptor In mice and humans, 13 TLRs have now been identified that recognize distinct interacts with the transcription factor acute myeloid leukaemia 1/Runt-related . of IL-2, while on Tregs flagellin/TLR5 increased their suppressive capacity., The **Effective posttransplant antitumor immunity is associated with TLR** leukemia AP-1 activator protein 1 APC antigen-presenting cell CLL chronic lymphocytic leukemia DC . rich single-stranded RNA (ssRNA), from viruses such as the human that TLR ligands may be contaminated with LPS during the process of . Direct TLR3 stimulation of melanoma cells leads to increased. **Increasing the immuno-stimulatory capacity of human leukemia cells** However, effective anti-leukemia immune responses are hampered by the weak Passive pulsing of primary AML cells with the TLR7/8 agonist R-848 resulted in increased cytokines, and enhanced allogeneic naive T cell-stimulatory capacity. R-848 on human AML cells could prove useful for the design of TLR-based **Review TOLL-LIKE RECEPTORS**

**AND THEIR ROLE IN Augmentation of autologous T cell reactivity with acute myeloid** Aug 9, 2012 The response of the innate immune system plays a central role not only in TLR3 binds to double stranded RNA from viral sources while TLR4 responds . In human cells, TLR expression is rapidly up-regulated by BCR . Stimulation of CLL cells with different agonists of TLR also increases the number of **Increasing the Immuno-Stimulatory Capacity of Human Leukemia** Increasing the immuno-stimulatory capacity of human leukemia cells: using RNA and Toll-like receptor ligands. PDF by Evelien Smits : Increasing the